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OM nucleic - nucleic search, using sw model

April 24, 2002, 03 25 22 ; Sour h Time 240,08 Seconds Run on:

(without alignments) 4895.125 Million cell updates/sec

US-09-525-998A-1 1368 Title: Periect score:

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IDENTITY\_NUC Scoring table:

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1861242 Total number of hits satisfying chosen parameters:

94002] SPGS, 428882819 PASIGNAS

Searched.

Minimum DB seq length: 0 Maximum DB seq length: 200000000

Post-processing: Minimum Match 0% Maximum Match 100

Maximum Match 100% Listing first 45 summaries

Database

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score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution. is the number of results predicted by chance to have a Pred. No.

/SIDS2/gegdata/geneseg/genesegn/NA2001.DAT.\*

					SUMMARIES	
		æ				
Result						
No.	Score		Match Length DB	DB	QI	Description
	1366.4	6.66	1368	14	AAQ49932	Lambda-derived FNF
5	1366.4	99.9		5	AAA95105	Human TNFR1 coding
m	1366.4			1.2	AAQ10883	30kD TNF inhibitor
4	1366 4	<b>৫ ৫</b>	2088	C1	AACR3946	Human 30 kDa TNF i
ďэ	1366 4			C1 C.	AA269170	Human tamour secre
ve:	1366.4			21	AAZ48475	Human tumour necro
7	1364.8			12	AAQ10955	Encodes human 55kD
30	1364.8			16	AA090513	p55 TNF-R gene. H
9	1363.2			7	AA020973	TNF-alpha binding
10	1363 2			7	AAQ24440	Encodes INF alpha
11	1363 2			CI	AA012215	Type I TNF recent

## ALIGNMENTS

AAQ4992 studing; cowa to mRNA; 1368 BP. 29-APP-1994 (first entry) Lambda derived INF R cDNA. AAQ49932; AAQ49932

Human, tumour prorosis factor receptor: TNF-R; interletkin: I receptor: II by fusion prof. i. There, IRE, ...; Ja hoxia, crot.us manaria; Thoumatoid attinitis, diabetes, munitipe solerosis; Septic shock; pulmenary fibrusis; Silicosis; allograft; xencgraft; rejection; qualti-nerges hit disease, cepsis, Inthamashou, alleriy; autoimmune dystunction; ss.

Homo sapiens. Lambda-gt10-7-etnfbp.

Logation/Qualitiers 1..1366 /\*tag- a /product- hINF-R 121..1363 ,\*taq- ∈ /\*tag- b /\*tag-120 W09319777-A sig\_peptide aut\_peptide 

14-00T-1993.

33W USC2938. 26 MAR-1993,

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therapy, diagnosis and assays for conditions mediated by TNF or IL-1, particularly in conditions in which both TNF and IL-1 play a causitive role. They may be used to treat cachexia, rheumatoid arthritis, diabetes, multiple sclerosis, pulmonary librosis and silicosis, cerebrai malaia, alivotail and actionally tejection in qualt verses host disease, sepsis, septic shock, inflammation, allergies and
                                                                                                                                                                                                                                                                                                                                               The sequences given in AAQ49931-32 encode human tumour necrosis factor receptor (TNF-R) and the sequences in AAQ49933-34 encode human interleukin I receptor (IL-IR). These sequences were used in the
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Ser, Thr and Ala. These linkers separate the individual mostics by such a distance that each component of the lusion protien is capable of Inding into the sevendary or terriary structure required for its biological activity. These fusion proteins may be used in
                                                                                                                                                                                                                  New Lusion protein tumour necrosis factor and human interleukin-1 receptor useful in therapy, diagnosis and assays of e.g. rheumatoid arthritis, diabetes, cerebial malaria, sepsis, etc.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            the linker may comprise 5:100 amino acids selected from Gly, Asp,
                                                                                                                                                                                                                                                                                                                                                                                                                    production of a fusion protein which conformed to one of the
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Sequence 1368 RP: 292 A: 424 G: 376 G: 276 T. 0 other,
                                                                                                                                                                                                                                                                                                                                                                                                                                                              TNF-R-linker-TNF-R-linker-IL-1R
IL-1R-linker-TNF-R-linker-TNF-R or
                                                                                                                                                                                                                                                                                                       Disclosure; Page 57–59; 85pp; English.
                    3120380 8226
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          TNF-R-linker-TNF-R
                                                              (IMMV ) IMMUNEX CORP
                                                                                                                                                   WPI; 1993-336592/42.
                                                                                                                                                                           P-PSDB; AAR42059.
                    30 - MAR-1942;
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181 aagtgochacaaaggaacctacttgtacaatgactgtccagggcrcgggggcaggatacggac 240 61 ggaatatacccctcaggggttattggactggtccctcacctaggggacagggagaagaga 120 121 gatagigidiceceaaggaaaatatateeaceteaaaataatilgeigiaee 180 12] gatagtgtgtgtccccaaygaaaatatatccacctcaaaataattcyatttyctytace 180 241 tgcagggagtgtgagagcygctcctfcaccgcttcagaaaccacctcagacactgcctc 300 agnityntriaaatgingaaaggaaatgggtnaggtggagatnitittignanagtggan 360 301 agetgetecaaatgeegaaaggaaatgggteaggtggagetetettgeacagtggae 360 461 egggacaccgtgfgtgtgtgcaggaagaaccagtaccggcattaltggagtgaaaacctt 420 461 Pagamamanyiyiyiyiyiyiyaqqaaqaammagtanegqmattattggagtgaaaaeett 420 12) ticcaqtgettelattgeagerletgesteaatyggagegtgeaerteteetgeeaggag 480 0; Gaps 1 algageoficionacegiaenigaeofigenegotaetaetaetaetaeteetegaeetatlagig 60 99.9%; Score 1366.4; DB 14; Length 1368; 1; Indels Mismatches Š Prod. 5 5 5 5 5 Conservative Similarity Matches 1367; Query Match 301 <u>a</u> ŝ 5 qq d C

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Human lNFRi coding sequence. 12-JAN-2001 (first entry)

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Polynucleotides comprising polymorphic variants of a reference sequence for tumour necrosis factor receptor 1 (INFR1), useful for studying the biological function of INFR1 and identifying drugs targeting the
                   INFR1: tumour necrosis factor receptor: polymorphism: human;
                                      tumour; cancer; apoptosis; bacterial infection; ds.
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                                                                                                                                     The present invention relates to polymorphic variants of the tumoun necrosis lactor receptor 1 (RNRA) gene. In present sequence is the colding applicate of the TMERI agene. The present sequence of the whole gene is given in AAA95102, AAA95103 and AAA95104. The polymorphisms were identified by amp liying and sequencing regions of the quee, Iwelyo polymorphic local word discovered, of these twive polymorphisms, region cause a change in the INFR protein. The infel polymorphisms may be useful for studying the biological tunction of INFR as well as for
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                                                                                                                                                                                                                                                           gene can be inserted into expression vectors for prepn. of INF inhibitor for use in the treatment of inflammatory and degenerative
                                                                                                                                                                                                                                        The whole
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          The sequence encodes the entire and kD TNF inhibitor. The cloue which the sequence was obtd, was isolated from a cDNA library prepd. From RNA form U937 cells treated with PMA/PHA. The wh
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 1; Indels
                                                                                                                                                                                                                                                                                                                                                                               Sequence 1988 BP: 439 A: 526 C: 578 G: 445 T: 0 other:
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                                                                                                                                                                                                                                                                                                                                See also AAQ10878, AAQ10884 and AAQ10907.
                                                                                           and -beta, useful as therapeutic agent.
                                                                                                                                       Disclosure, Fig 21, 142pp, English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                           99.98;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Sest Local Similarity 99.9%
Matches 1367; Conservative
WPI; 1991-073847/11.
                      P-PSDB; AAR10986.
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889 incaagototaticattittitijiaaalojadaoolgaaaagagagagagagatigaa 948
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        inhibitor; antiinflammatory, Tumour Necrosis Pactor, interleukin,
                           cocaccitygycttcaytcocyfycccayttcoaccttcacctorageteeacctatace
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89US-0381090.
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18-JUL-1989;
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Hale KK, Brewer MT, Thompson RC;

King MW,

Squires C,

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The present invention relates to Tumbour Newtonis Factor (TMF) inhibitors (see ARR376F) and ARR376FS), which have NFF inhibitory activity, the nevel TMF inhibition of the present in-centur are set, to their apentic agents for inhibiting the activity of TMF and interleukin (IL-1), and for resulting inflammatory, and degenerate activity of TMF and interleukin (IL-1), and present segmence is the centure of the present segmence in the inhibiting our inhibitors of 36 kDm inhibitors in the tenture of 36 kDm inhibitors.
                                                                             Novel 30 kDa tumor necrosis factor inhibitor analog comprising a for native systeing tesidus cross-linked with polychylene glycol, useful for treating inflammatory and degenerative diseases mediated by
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Tumour necrosis factor binding protein, TNF, insoluble protein; agonist; anti-inflammatory; antimalarial; treatment; septic sheek; inflammation; autoimmune glomerulonephritis; cerebral malaria; immune response;
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their (in)soluble fragments (Ia) and their saits that can bind tumour necrosis factor (TNF) The products of the invention have anti-inflammatory and antimalarial activity. (I) and (Ia) are used (i) to treat diseases in which TNF is involved (e.g. septic shock, autoimmune glomeruloucphritis, cerebral malaria, immune responses and inflammation), (ii) to purity TNF, (iii) to identify TNF (an)sagonists and (iv) for diaphystic determination of TNF in body fluids. Antibodies raised against (I) are used for affinity puritication of (I). This sequence encodes a tumour necrosis factor binding protein described in the method of

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                                                                                                                                                                                                                                                                                                                                                                 New insoluble proteins, and fragments, that bind to tumor necrosis factor, used to treat e.g. septic shock or cerebral malaria
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                                                                                                                                                                                                   (HOFF ) HOFFMANN LA ROCHE & CO AG F.
                                                                                                                                                                                                                                                                                                                                                                                                                               Claim 4; Fig 1; 25pp; German.
                                                                              90EP-0116707.
                                                                                                                                        89CH-0003319.
                                                                                                                                                               9UCH-UUUU746.
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                                                                                                                                                                                                                                            Brockhaus M,
                                                                                31-AUG-1990;
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EP939121-A2
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                                                                                                                                                                                                                                                                   Schlaeger
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The invention provides antisense compounds targeted to human tumour necrosis tactor receptor type 1 (TRES) RNA. These antisense compounds can be used in a southed of lithilitims the expression of TRES bunch of its or tissues. The antisense compounds specifically hybridize with one or tasues, the antisense compounds specifically hybridize with one or about motions of nucleic acid molecules encoding TRES, ultimately modulating the function of nucleic produced. The antisense compounds and method are useful as research reagents and diamnostics, and in the treatment and prophylaxis of information of uncertions is information of uncertions. The present sequence represents the uncleotide sequence of bunch INES (Garhaux Acon No. NSSNI).
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99.9%; Pred. Ko. 9;
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intlammation; tumour formation; INFR; anticancer; its
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Recombinant constructs are used to transform cells to confer
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Expression of this receptor fragment. The p55 INF R can be shot in response to different indo ind adents, to a photon approach to the receptor fragment. The p55 INF R can be shot in response to different indo ind adents, to a photon of the receptor whose structure after the receptor fragment. The receptor of the receptor whose structure extracellular domain. This region is located object to a site of clearage of the region of the encoded protein was used to create the chimeras between human p55 INF R and murrain or pidential grade artor receptor (EGF-R) that are represented by AAR35072-17 and substitutions are subjected to detection murrains (AAR35072-17) and substitutions. ARR35073-17 this spacer region distributions with series of the spacer region of the Spacer region and the Spacer region of the Spacer region are represented by AAR35077-11. This spacer region are subjected to detect on detection murrains (AAR35077-11) and substitutions. ARR3507-11. inhibiter of a protease that is capable of Freather the soluble INF-R from the cell bound INF-K. Fragments of these inhibitors can be seen in AAP75017-9, AAP75025, AAP75043-5 and AAP75047-4. These protease important of these. The shedding of the receptor is independent of the side chain identity of those residues, with the exception of a limited dependence on the identity of Val 173. Mutations which alter the This sequence represents human ps; tumour necrosis factor (INF R) DNA charloimation of the protein adversely effect the shedding process. I mutations shown in AAR75013-47 were introduced in order to create an pSS: Fumour neriosis factor receptor: INE-E; Leman, mainner edimera; epidermal growth factor receptor: EdE-E; protease; inhibitor; these professe New protease capable of cleaving soluble tumour mecrosis factor (INF) receptor - from cell-bound IMF- receptor, useful for antagonising deleterious effects of INF. inhibitors can be used for enhancing INF function. "possible poly-A signal" Batkie M. Bruketisch C. Varfolomeer E. /\*\*.aq= a /product= p55 TNF-R photbol myristate acetate, PMA, ss. Disclosure: Fiq 1: 40pp: English. AAQ90513 standard; DNA; 2175 BP (YEDA ) YEDA RES & DEV CO LID. 9-01-0107268. 3460 0075742. 2144...2149 19-3AN-1996 (first entry) 256...1623 /\*t.aa-WP1; 1995-194342/26. P PSDB; AAR75084 p55 INE-R gene. Homo saplens. 12-00T 1993; 11 001 1934, misc\_siqual AUG175742-A 04-MAY-1995 AAU90513: 

edec :

Indels

99.8%; Score 1364.8; DB 16; Length 2175;

Pred. No. 0:

99.98;

Matches 1356; Compercative

Best Local Similarity

Ouery Match

Sequence 2175 BP: 474 A; 642 C; 623 G; 456 T; P other;

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/note- "encodes the extracellular domain of human
TNF alpha receptor"
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binding protein which was obtd. from a human placental cDNA library of in lambda gtil using a prate (AAQ20974). The DNA place provides the contractional domain of human INP alpha receptor and as such it is used; for treating discases where TNP alpha is irrelyed as such it is consative agent. e.g. cachexia, sepsis and autoimmune discases, consettically rheumatoid arthritis. See also AAQ20074

Sq. specifically rheumatoid arthritis. See also AAQ20077

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library in 9110 was screened with probe AAQ29236. Ten hybridising clones were plague purified and ThNA size determined by TAGE against an ECO K of greated phage DNA. The inserts of two CDNA clones were then sequenced the coding region of the majority of the human TNF alpha sequenced the coding region of the majority of the human TNF alpha SFMD receptor was isolated as an ECORI fragment encoding 374 amino acids, and planta into a mammalian cell extression vector, resulting in pITNFR. A derivative of the TNF-alpha receptor was produced by
                                                                                                                                                                                                                                                                                                                                                                                                            This sequence encodes human TNF-alpha 55kD receptor . A placenta cDNA
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DNA sequencing confirmed this contained the designed DNA sequence.
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treating autoimmann disease, scylic shock, HTV etc.
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                                                                                                                                                            Feldmann M, Gray PW, Turner MJC;
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                                                                                                              (CHAR-) CHARING CROSS SUNLEY RES CENT
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                     91WO-GB01826.
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Matches 1365, Conservative
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481 aaacagaacaccgiqtqcacctgccatgcaggtititititicaayagaaaaacggiqtgtu
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              Location/Qualifiers
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type inmost Necrosis Factor Binding Protein i is the solubic form of type i INF receptor and constitutes a fragment of the cell surface form of this receptor, corresp. to its extracellular domain. There is no characteristic poly(A) addition signal near the 3° end of the cENA. The sequence ACTAAA (tag m) may serve as an

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LambdaTNF-R2 encodes the complete human TNF-R2 and was used to
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construct a plasmid (pADTNF-R) expressing the product the same way as pADTNF HP (see AAQU6282). See also AAQU6282-Q06285.
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                                                                             Sequence 2141 BP; 455 A; 633 C; 593 G; 460 T; 0 other;
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Best Local Similarity 99.74
Matches 1364; Conservative
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1113 eccqqtgaetgteccaaett/gegyeteccegeayagagqtggeaceaeeetateagggg 1172
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                                                                                     Modulating artivity of tumour necrosis factor receptor - using peptide(s), antibodies, etc. which interact with relited regions of receptor or effector protein, for controlling auto-immune
                                                                                                                                         p55 Tammar necrosic factor receptor
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256..1623
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99,6%, Pred. No. 8.2e 315;
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                                                                                                                                   1216 gotgannnatonttgnganagnnntngnntnngsnnnnatonnnaapppppttpagaag 1278
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841 eccaecrigggettragtecegigecrossiteraretteaeqiecagetecaectataee 900
                                                    90] eccyatyaectgtermaantttgeggeteeeeqmaqagaggtggeameaeeetateagggg
                                                                                                                                                            Plasmid Tumour Necrosis Factor-Binding Protein 15 cDNA insert.
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Himmler A, Maurer-Fogy I,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Location/Qualifiers
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             AAQ06282 standard; DNA; 1334 BF
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 633 trecagigeticaaligeageeleigeeleaalgagaeegigeaeeleeteelgeeaggaa 692
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                                                                                                                                      from induced TNP-induced fibrosarcoma cells. A TNP-BP had been isolated from the urine of patients with uraemia and probes/primers were constructed from the determined union acid sequence.

To produce a vector expressing a solutio form of TNP-binding protein, this plasmid was cut with Xmml, amplified by PCR and the
                                                                                                                       pPNF-BP15 is one of 30 positives clones in a screened cDNA library
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   gataqtgigtgiceeccaaggaaaatatateeareeteaaaataatilogattilgetgiee
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               DNA encoding TNF binding protein and TNF- receptor - used in tumour treatment and to understand mechanismsm to TNF action
                                                                                                                                                                                                                                                                                                                                    PAD-CMV1 (AAQ06283) to give the required plasmid pADTNF BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Indels
                                                                                                                                                                                                                                                                                                                                                                                                Sequence 1834 BP; 299 A; 409 C; 342 C; 284 T; 0 other;
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                                                                                                                                                                                                                                                amplified DNA cut with BamH1 and EcoR1.
                                                                               Disclosure; Fig 1(1-3); 51pp; German.
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                                                                                                                                                                                                                                                                                                    her common grant may more granted and the accepted at the acceptation 1112
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Rat Tumour Necrosis Factor-Receptor 8 cDNA.
                                Disclosure: Fig A(1-2); 51pp; German.
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